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Docket No.: 202443US0

## IN THE UNITED STATES PATENT & TRADEMARK OFFICE

IN RE APPLICATION OF

Veronique CHEVALIER, et al.

**EXAMINER: M. WILLIS** 

SERIAL NO: 09/784,179

FILED: FEBRUARY 16, 2001

GROUP ART UNIT: 1619

FOR: COMPOSITION BASED ON

N-CHOLESTERYLOXYCARBONYI\_ 4-PARA-AMINOPHENOL AND

HYDROQUINONE OR ONE OF ITS DERIVATIVES

## DECLARATION UNDER 37 C.F.R. 1.132

ASSISTANT COMMISSIONER FOR PATENTS WASHINGTON, D.C. 20231

SIR:

- I, Dang Man PHAM, hereby declare:
- 1. I am employed by L'ORÉAL as an engineer and have experience with skin depigmenting and/or lightening agents.
- The following observations and experiments were carried out by me or under my direct supervision and control.
- 3. The depigmentation activity of combinations of compounds was determined using the techniques set forth in Example 1 of the present application. In accordance with the claimed invention, the depigmentation activity of N-cholesteryloxycarbonyl-4-para-aminophenol (6.77 x 10<sup>-5</sup>) and hydroquinone (8 x 10<sup>-5</sup> M), alone and in combination, was determined. These results are set forth below.

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	% inhibition observed	% inhibition theoretical
Hydroquinone (8 x 10 <sup>-5</sup> M)	25	
N-cholesteryloxyearbonyl-4-para- aminophenol (6.77 x 10 <sup>-3</sup>	No detection	
Ilydroquinone (8 x 10 <sup>-5</sup> M) + N-cholesteryloxycarbonyl-4-para- aminophenol (6.77 x 10 <sup>-5</sup> M)	31.8	24.4

For purposes of comparison, the depigmentation activity of N-ethyloxycarbonyl-4paranminophenol (6.77  $\times$  10<sup>-5</sup> M) and hydroquinone (8  $\times$  10<sup>-5</sup> M), alone and in combination, was determined. These results are set forth below.

Hydroquinone (8 x 10 <sup>-5</sup> M)	% inhibition observed 26.12	% inhibition theoretical
N-ethyloxycarbonyl-4-para- aminophenol (6.77 x 10 <sup>-5</sup> M) Hydroquinone (8 x 10 <sup>-5</sup> M) +	13.64	
N-ethyloxycarbonyl-4-para- aminophenol (6.77 x 10 <sup>-3</sup> M)	16.48	39.76

Thus, both experiments were performed using the same concentration of hydroquinone compound (8 x  $10^{-5}$  M) and of aminophenol compound (6.77 x  $10^{-8}$  M).

- 4. As shown above, the claimed combination of hydroquinone and Ncholesteryloxycarbonyl-4-para-aminophenol resulted in synergistic inhibition activity (31.8% actual inhibition vs. 24.4% theoretical). In contrast, the combination of hydroquinone and Nethyloxycarbonyl-4-paraaminophenol did not result in synergistic inhibition activity and, in fact, resulted in less than additive activity (16.48% actual inhibition vs. 39.76% theoretical).
- 5. Given that the same concentration of aminophenol compound was used in both experiments, the difference in inhibition activity between the claimed hydroquinone/aminophenol combination and the comparative combination was unexpected

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and surprising, and demonstrates the improved depigmentating activity of the synergistic combinations of the present invention.

- 6. The undersigned petitioner declares further that all statements made herein of his own knowledge are true and that all statements made on information and belief are believe to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of this application or any patent issuing thereon.
  - 7. Further deponent sayeth not.

April 18th 2003